# Avoiding Misdiagnosis in Patients with Neurological Emergencies

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Avoiding Misdiagnosis in Patients with Neurological Emergencies

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Approximately 5% of patients presenting to emergency departments have neurological symptoms. The most common symptoms or diagnoses include headache, dizziness, back pain, weakness, and seizure disorder. Little is known about the actual misdiagnosis of these patients, which can have disastrous consequences for both the patients and the physicians. This paper reviews the existing literature about the misdiagnosis of neurological emergencies and analyzes the reason behind the misdiagnosis by specific presenting complaint. Our goal is to help emergency physicians and other providers reduce diagnostic error, understand how these errors are made, and improve patient care.

1. Introduction

Approximately 5% of emergency department (ED) patients present with neurological symptoms [1]. The most common symptoms or diagnoses that these patients have are headache, dizziness, back pain, weakness, and seizure disorders [2–6]. In recent years, improved time-dependent treatments for patients with acute neurological emergencies have been developed, increasing the importance of a rapid and accurate diagnosis. Underdiagnosis may have disastrous consequences. Conversely, overtesting leads to inefficient resource utilization that is undesirable for both economic and medical reasons.

2. Methods

A PubMed search on February 8, 2012 for the intersection of “misdiagnosis” and “neurological emergency” as title or abstract words resulted in 88 results. In addition to this literature review, we will incorporate experience from over 30 years of ED clinical practice, teaching medical students, and residents, over a decade of evaluation of medicolegal cases, and analyzing diagnostic errors committed by our colleagues and ourselves in peer review.

We review the existing literature about misdiagnosis of nontraumatic neurological emergencies in general, and then by specific presenting complaints. We conclude by analyzing the reasons for misdiagnosis. Our goals are to help emergency physicians (EPs) and other front-line clinicians reduce misdiagnosis of patients with neurological emergencies and to be hypothesis generating so that we can better study and understand misdiagnosis in these patients and to improve patients’ clinical outcomes.

3. Results

3.1. General Studies about Misdiagnosis of Neurological Emergencies

Few high-quality data on the subject of ED misdiagnosis of patients with neurological emergencies exist. Most papers on misdiagnosis of patients with neurological emergencies focus on patients with a particular diagnosis or presenting symptom. Only a few analyze the general topic of all-comers with neurological symptoms [2–4]. There are methodological problems with all of these articles. The EP’s diagnosis is made earlier in a patient’s course. Therefore, less historical information is usually available, the natural course of the disease process is less well defined, and almost always, fewer results of diagnostic testing are available. The primary job of the EP is to ensure clinical stability and proper disposition of a patient, both of which are possible without necessarily making a specific etiologic diagnosis. Therefore, EP’s charted diagnosis is often a tentative one, or even simply
Two other studies did not so much compare EP versus neurologists’ diagnoses, as categorize the types of ED neurological emergencies [2, 5]. In all studies of emergency neurological consultations, stroke, headache disorders, seizures, and dizziness make up a large majority of the patients [2–5, 7]. In Hansen’s study, which analyzed 500 neurology consultations at a tertiary U.S. academic hospital, 4.8% of total ED patients had a neurology consultation (1/3 of the number in the French study). The mean length of stay for those patients was 7.4 hours (significantly longer than for the average ED patient—4.9 hours) and remarkably similar to the “just under 8 hours” in the Canadian study. In the latter study, it is interesting to note that patients with diagnostic ambiguity stayed in the ED much longer than those where there was either agreement or disagreement (between consulter and consultant) about the diagnosis.

Although all these studies have limitations, there is a common theme that runs through them. Diagnosis of patients with neurological emergencies is imperfect. There is significant underdiagnosis (which threatens patient safety) and overtesting (which wastes resources). Patients with stroke, dizziness, headache, and seizures are the most common sources for these errors. In a study of unplanned ED return visits, many of which were due to missed diagnoses, headache and vertigo were among the most common presenting symptoms [8]. Apart from the studies discussed above, most others have analyzed misdiagnosis by either specific presenting symptoms (e.g., headache) or by specific diagnosis (e.g., SAH).

### 3.2. Headache

Headache accounts for roughly 2% of ED visits, of which only a very small percentage have serious secondary causes [9]. This “needle in the haystack” phenomenon may lead clinicians to not consider serious secondary causes. Deciding which patients to investigate beyond clinical evaluation can be difficult; history and physical examination must focus on uncovering “red flags” that suggest the need for further testing [10].

Much of the literature about misdiagnosis of headache focuses on subarachnoid hemorrhage (SAH) [11–13]. While older literature showed a misdiagnosis rate from 12–25%, the latest data based on misdiagnosis restricted to the ED puts that figure at 5% [13]. Recurring reasons for misdiagnosis include not considering the full spectrum of presentations, not following an algorithmic workup and not understanding the limitations of the tests in that workup [11, 12].

Regarding presentation, not all patients with SAH have a truly abrupt onset of their headache [14]. In some patients, the headache improves after analgesics including triptans [15]. Some patients present with prominent vomiting or fever and neck pain or with hypertension, each of which can divert the physician’s diagnosis to other less serious problems such as gastroenteritis, viral syndrome, or hypertensive crisis [11, 12]. Patients with SAH do not necessarily “look ill,” have any neurological deficits or meningism.

Even in the less acuity-skewed population of a neurology practice, some argue for a lower threshold for imaging in patients with new-onset headache [16]. This is probably
3. Dizziness. As with headache, dizziness has both benign and serious causes that can be difficult to distinguish from one another. Diagnosis of the dizzy patient is inherently fraught with problems. The diagnostic algorithms that doctors are taught may be flawed. Increasing evidence suggests that the traditional “symptom quality” approach (“what do you mean by “dizzy”?”) is less effective that a new “timing and triggers” approach, in which the physician asks about the temporal characteristics of the symptoms [26-28]. Data suggests that whether the patient uses the word “vertigo” or “spinning” versus “dizzy” or “lightheaded” is not so useful in determining etiology. “Vertigo” versus nonspecific dizziness does not help predict etiology in dizzy patients [29, 30]. For example, patients with BPPV often use nonspecific (non-vertigo) descriptors for their symptoms [31] and patients with clear-cut cardiac causes of dizziness often complain of “vertigo” [32].

Because of the prevailing paradigm, EPs may have an overly generalized approach to dizzy patients [28]. There is also a significant overlap between the presentations of benign (vestibular neuritis and labyrinthitis) from serious (cerebellar and brainstem stroke) presentations [33, 34]. Deficits in physician knowledge may also contribute, for example, documentation of nystagmus is often inaccurate [35]. Finally, lack of understanding of the limitations of neuroimaging is another issue. Some EPs incorrectly believe in the sensitivity of CT to exclude posterior circulation stroke [28], which may also be undetected by MRI in the first 48 hours [33].

Neurologists may also have difficulty diagnosing dizzy patients in the ED. Royl reported on 475 patients seen in a German neurology ED staffed by neurologists. Of the 124 patients for whom followup was available, 43% of ED diagnoses were “corrected” [36]. Six percent of the patients diagnosed with benign conditions were changed to serious ones and 23% of the serious ones were reversed to benign. In a California study of ED patients discharged with an ICD-9 code compatible with dizziness, there was an increase in the incidence of adverse cerebrovascular events in the next 30 days, suggesting that an important diagnosis had been missed [37].

The most feared misdiagnosis of dizzy patients is stroke. These are usually ischemic strokes of the brainstem and cerebellum. In one series of 240 consecutive cerebellar strokes, 10% presented as an acute vestibular syndrome (AVS) suggesting a peripheral cause [34]. Nearly all of these patients had posterior inferior cerebellar artery strokes. Patients with misdiagnosis may have poor outcomes due to posterior fossa edema and brainstem compression [38]. Distinguishing stroke from benign peripheral causes is critical, not just to treat the acute complications, but also to evaluate and treat the underlying vascular lesion in order to prevent a second event [39].

3.4. Back Pain. Along with headache and dizziness, back pain is very common and most patients have benign, self-limited causes. With back pain, there are fewer “needles” in a larger “haystack.” Common causes of cord or cauda equina compression include herniated disk, tumor, abscess, and hematoma. In primary care practices, all four of these etiologies amount to roughly 1% of patients with back pain [40]. Diagnoses generally require MRI, thus setting up the classic tension between resource utilization versus patient outcomes [40]. Surprisingly, few data exist about prevalence in the ED, although it is likely higher due to skewed acuity. Red flags include new pain in patients > age 50 years, a history of cancer, fever, weight loss, an immunocompromised state, intravenous drug use, recent bacteremia or urinary tract infection, pain that is worse with rest or at night, sphincter symptoms, bilateral sciatica, failure to improve over weeks, anticoagulation, and recent spinal procedure [41]. Some patients have no identifiable red flags.

Cauda equina syndrome (CES) can be misdiagnosed and/or lead to malpractice claims because of inadequate history, physical examination, or communication between physicians and between physicians and nurses [42]. In one small series of 32 patients with CES, fewer than 20% presented with the classic presentation of bilateral sciatica, leg weakness, saddle anesthesia, and sphincter dysfunction [43]. The most common reason for misdiagnosis in that series was failure to consider the diagnosis. In a retrospective study of 23 patients with suspected CES, the diagnostic accuracy of individual findings of urinary retention, frequency, incontinence, or altered urinary or perineal sensation ranged from 57–65 percent [44]. In another retrospective study of 58 consecutive patients of suspected CES, having 2 of the following 3 findings (bilateral sciatica, subjective urinary retention, or rectal symptoms) increased the likelihood of a positive MRI 48 folds [45].

Another finding that is often ignored is ataxia or new frequent falls. In a study of 63 patients with nontraumatic cord
compression or CES, nearly one in four patients had ataxia or gait difficulty with neither sensory nor motor findings [46]. For spinal epidural abscess (SEA), the incidence of the typical triad of back pain, fever, and neurological deficit is low [47–49]. For all these reasons, misdiagnosis or delayed diagnosis is common [47, 50, 51].

Some algorithms include measuring inflammatory markers such as the erythrocyte sedimentation rate (ESR) or C-reactive protein (CRP). Sensitivity for an ESR (>20 mm/hour) in infectious causes of cord or cauda equina compression such as SEA or vertebral osteomyelitis, range from 76–95%; the corresponding figures for CRP are 82–98% [41, 52, 53]. For neoplastic causes, the sensitivity of ESR (>20 mm/hour) is 78% [54]. For the ESR, increasing the threshold value increases specificity at the cost of sensitivity.

MRI with gadolinium is the test of choice for most of these problems [50, 55]. A significant diagnostic issue is that in many practice settings, obtaining an urgent MRI can be difficult or impossible. In the absence of a strict diagnostic algorithm that forces the clinician to get the MRI, even if that requires a transfer, the lack of availability of the diagnostic gold standard may lead to misdiagnosis. Both SEA and epidural tumor often affect multiple areas of the spine; therefore, it is important to decide which part of the spine to image. Some experts recommend that the entire spine should be imaged [55]. Finally, there are issues of resource utilization; in one study of 106 patients undergoing MRI for possible SEA, only 7 were positive [56].

3.5. Weakness. The majority of patients presenting to an ED with generalized weakness have a variety of “toxic-metabolic” problems including electrolyte abnormalities and dehydration, medication side effects, and systemic infections. Acute neurological causes of generalized weakness include uncommon diagnoses such as Guillain-Barré syndrome (GBS), transverse myelitis, myasthenia gravis, and periodic paralysis, as well as rarer conditions such as tick paralysis, botulism, and others.

Misdiagnosis of GBS is common [57, 58]. In a series of 20 ED cases, most patients complained of weakness but some presented with paresthesia [57]. 15 of the 20 patients were incorrectly diagnosed on their first ED visit. Four patients initially presented with sensory symptoms. Six of the 20 had normal cerebrospinal fluid (CSF). The notion that the CSF protein is always elevated is wrong; normal CSF protein is common in the first week of GBS [59]. Misdiagnosis of Lambert-Eaton syndrome is also common [60]. Similarly with myasthenia gravis (MG), the mean time to correct diagnosis is over a year [61]. The variability of specific symptoms and their timing makes diagnosing MG difficult. Therefore, to diagnose MG in the ED, it is essential to know the full spectrum of possible presentations [62]. In transverse myelitis, asymmetric cord involvement can lead to atypical presentations that make diagnosis more difficult [63]. Unless one checks the serum potassium at the time of symptoms, periodic paralysis may be missed. In very rare conditions such as botulism and tick paralysis, initial misdiagnosis is quite common [64, 65].

Serious misdiagnosis can occur in patients with stroke and transient cerebral ischemia (TIA) who present with focal weakness. Various studies have reported that the rate of ED misdiagnosis of stroke varies widely, from as low as 2% to as high as 56% [66–72]. To some extent, the variation relates to study design. The two most recent North American studies found rates of misdiagnosis of roughly 10% [69, 70]. Current diagnostic scoring systems for ischemic stroke emphasize lateralizing motor findings [72–74]. Factors associated with stroke misdiagnosis include young age, posterior circulation or sensory symptoms, and lack of lateralizing weakness [38, 75, 76]. Some stroke patients have NIH stroke scores of zero [77]. Many of the patients in that study had posterior circulation strokes. Finally, physicians must understand that some strokes, even of the anterior circulation, present with atypical symptoms like neuropsychiatric symptoms or abnormal movements at stroke onset [78].

Diagnosis of TIA is more difficult because most TIA patients are neurologically intact by the time they are in the ED. An early study found a misdiagnosis rate by the emergency physician of 6% [79]. A major limitation of this study is that neurologists made their diagnosis by reviewing the ED chart, not by independent clinical evaluation. Another study found a misdiagnosis rate by EPs of 60%, with factors leading to misdiagnosis including gradual onset of symptoms, prior similar episodes, and nonspecific symptoms [80]. The most recent and methodologically sound study found an ED misdiagnosis rate of 36% [81]. It also showed that the presence of headache, involuntary movements, and dizziness were all associated with a non-TIA diagnosis.

Apart from the aforementioned studies of SAH, there has not been much systematic study of misdiagnosis of intracerebral hemorrhage (ICH). Both CT and MRI are very sensitive for ICH; therefore, when physicians perform brain imaging in patients with weakness, they will find ICH when it exists. Although ICH often presents more dramatically than ischemic stroke, there is significant overlap. It is certainly possible that patients who are not imaged will be misdiagnosed.

3.6. Seizures. As with TIA, the diagnosis of a seizure often depends entirely upon the history of an event that the physician has not witnessed. Therefore, it is important to try to obtain information from any witnesses of the event, and to gather what data one can from the physical examination to distinguish the causes of these transient episodes of loss of consciousness. The most common issue is distinguishing syncope from seizure, but one must also separate true seizures from pseudoseizures (also referred to as psychogenic seizures and nonepileptic attack disorder). Of these three conditions, syncope is by far the most common.

In one review, the misdiagnosis rate overall for seizures in both children and adults ranged from 5–30% [82]. In adult patients incorrectly diagnosed with seizure, the most common final diagnoses were syncope and pseudoseizures [82, 83]. In children, various benign paroxysmal disorders such as breathing holding spells and night terrors were the most common final diagnoses [82]. The source of
misdiagnosis was not entirely from the ED. It is important to note that electroencephalography (EEG) is not specific for seizures and lacks sensitivity. That is to say, an abnormal EEG does not exclude pseudoseizures and a normal EEG does not exclude true seizures or confirm pseudoseizures [84–87].

There are several characteristics that help emergency clinicians to distinguish between syncope, epileptic seizures, and pseudoseizures. Syncope may have a prodromal sensation of warmth, lightheadedness, sweating, and facial pallor, and is often precipitated by various triggers. The event starts rapidly and recovery is prompt. In cardiac causes of syncope, palpitations or chest pain may occur together. Importantly, however, “convulsive syncope,” in which the faint is accompanied by some tonic-clonic jerking due to brain hypoperfusion is common [85, 88, 89]. Tongue biting may occur but it is usually at the tip of the tongue [90]. Urinary incontinence is unusual but may also occur [85, 86].

Patients with true seizures often have a preceding aura or repetitive movements (chewing or lip smacking), lateral biting of the tongue or cheek, facial cyanosis, sphincter incontinence, head turning towards one side, and postictal confusion that is slow to resolve [85, 90, 91]. Postevent neurological examination may show focal deficits [86]. Though not extensively studied, transient anion gap acidosis is also associated with true seizure [92]. Up to 36% of patients with “intractable seizures” actually have pseudoseizures [84]. Patients with pseudoseizures may show side-to-side head movements, changing symptoms if multiple spells, gradual onset and waxing and waning during the spell, rapid recovery, and bizarre movements involving the entire body without any “logical” march [86]. Tongue biting and incontinence are less common in patients with pseudoseizures compared with true seizures [90]. In many ED patients with transient loss of consciousness, a definite diagnosis will not be possible. Coordination of subsequent care for testing not available in an ED such as tilt-table testing, continuous loop ECG monitoring, or video-monitored 48-hour EEG testing may help to reduce misdiagnosis.

3.7. Conversion Reaction (Functional Neurological Symptoms).
Conversion reactions overlap the symptom-oriented discussion above. The most common conversion reactions relate to weakness and seizures [93]. One report of ED patients diagnosed with conversion reaction who later proved to have organic disease emphasizes that misdiagnosis often relates to patients having symptoms atypical for organic disease (e.g., “I’ve never seen anything like this before” or symptoms being “non-anatomic”) [94]. Hoover’s sign (weak hip extension that becomes normal on testing contralateral hip flexion) was found to be moderately sensitive and very specific for functional weakness [95]. In the specific setting of possible ischemic stroke, it is obviously preferable to not give thrombolytic therapy to someone who does not have a stroke. However, patients with stroke mimics who are CT negative, have never been reported to have hemorrhagic complications [96–98]. Although a high degree of diagnostic accuracy is possible [99], EPs should be very hesitant to make a diagnosis of conversion reaction in the ED.

4. Discussion
Before discussing the data, it is important to acknowledge their limitations. The literature does not contain high-quality data on this subject, and the data derived from our experience with quality assurance and medicolegal case review is by definition skewed towards cases with poor outcomes. We acknowledge that our conclusions are limited by the weakness of the data upon which they are built; however, we believe that this is the best available analysis of the data.

Misdiagnosis contributes to medical malpractice in the ED and patient harm [100, 101]. The underlying reasons included inadequate history and physical examination, failure to order and correctly interpret tests, and failure to obtain a consultation [101]. In Table 1, we have listed potential causes for misdiagnosis of patients with neurological emergencies. Researchers in the field of diagnostic error often characterize errors in terms of cognitive analysis, which is useful for research [102]. Herein, however, we will categorize reasons for error in everyday terms that average clinicians will not only understand but also relate to.

Time pressures, frequent interruptions, and distractions are common in the ED. For stroke, time pressures related to thrombolytic use force EPs to “diagnose” a stroke within minutes of the patient’s presentation, when key historical details may be unavailable. For the less common diagnoses, the “needle in the haystack” phenomenon exists. “Classic” triads and the “typical” symptoms that are emphasized in medical education are often absent. Preconceived notions are sometimes wrong. In addition, examining the nervous system is more complicated than examining the heart or lungs. Charting systems, designed to maximize billing, discourage good documentation. The best test for some conditions, MRI, is often unavailable. Incidental findings on physical exam or imaging tests may distract and prematurely stop the workup. A false normal study (due to interpretation error or imaging the wrong site or at the wrong time) may do the same.

Over testing can also result in patient harm. Incorrect diagnosis of a seizure often leads to anticonvulsant use or driving restrictions. With respect to investigations, ED use of CT more than tripled over the period 1995–2007 [103] and interestingly there is a 3-fold variation of CT use across individual physicians [104]. Apart from obvious issues of diagnostic accuracy, evidence is mounting about the long-term consequences of increasing radiation exposure [105]. Furthermore, incidental findings drive further investigations, which may lead to adverse consequences [106].

Finally, it must be acknowledged that some degree of misdiagnosis is unavoidable [107]. Making every diagnosis every time has costs. Even immediate ED neurological consultation will not lead to diagnostic perfection. Both Moeller and Royl showed that neurologic evaluation in the ED was still associated with some misdiagnosis [3, 36]. In
Table 1: Some reasons for misdiagnosis in patients with neurological emergencies.

<table>
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<tr>
<th>History</th>
<th>Headache</th>
<th>Dizziness</th>
<th>Back Pain</th>
<th>Weakness</th>
<th>Seizure</th>
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<tr>
<td>Patients use words like “migraine” and “sinus infection” that may mislead the physician. Beware previous diagnoses; they might be wrong.</td>
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<td>The use of the word “vertigo” versus other dizziness descriptors is not etiologically useful.</td>
<td>Patients use word “sciatica” which may lead physicians to diagnose sciatica.</td>
<td>Stroke patients may complain of “clumsiness” or “my arm felt like lead” rather than “weakness”.</td>
<td>Patient (or witness) says “seizure” after a faint Seizure patients often present after the seizure with only an altered mental status or with a postictal “Todd’s” paralysis</td>
</tr>
<tr>
<td>Physical exam</td>
<td>Patients with SAH may be well appearing and neurologically intact.</td>
<td>Patients with small posterior circulation strokes can mimic a peripheral vestibular presentation.</td>
<td>Patients with serious causes of back pain can present without neurological deficits.</td>
<td>Patients with stroke can present with just about any focal deficit depending upon the occluded vessel. Myasthenia patients’ symptoms wax and wane. GBS patients’ first symptoms may be purely sensory.</td>
<td>Patients may be lethargic, but neurologically intact.</td>
</tr>
<tr>
<td>Diagnostic testing</td>
<td>For SAH, CT sensitivity is good but decays with time. CT has poor sensitivity for CVST and dissection.</td>
<td>CT is a poor test for cerebellar and brainstem infarction</td>
<td>No MRI available; MRI must target the correct segment(s) of the spine.</td>
<td>False normal CT in early stroke</td>
<td>EEG often not available in the emergency department. Not performing LP in seizure patients who may have encephalitis or neurocysticercosis.</td>
</tr>
<tr>
<td>Preconceived notions</td>
<td>Headache improved with triptans so is not a serious secondary cause.</td>
<td>Posterior circulation strokes are obvious or devastating events</td>
<td>All patients with SEA have risk factors or fever, or neurological deficits</td>
<td>Young people do not get strokes</td>
<td>Seizures (or seizure-like movements) are sometimes seen with strokes. Convulsive movements are common in syncope.</td>
</tr>
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CVST: cerebral venous sinus thrombosis, SAH: subarachnoid hemorrhage, CT: CAT scan, MRI: magnetic resonance imaging, SEA: spinal epidural abscess.
a study of malpractice cases against neurologists, EPs were codefendants in 44% of cases [108]. These data corroborate the obvious conclusion that simply consulting a neurologist does not eliminate potential errors. Because some diagnostic uncertainty is inevitable, explicit communication between physicians, physicians and patients and thoughtful coordination of followup care after the ED phase become critically important.

5. Conclusions and Solutions

A full analysis of the reasons behind these potential errors and solutions to the problems is beyond the scope of this review. However, several generic issues exist. Less than 20% of emergency medicine residencies require a neurology rotation [109]. Education is mostly lecture based; however, many of these lessons are best taught by studying real-life patients at the bedside. Important as education is, diagnostic error is frequently not the fault of a misinformed individual. Numerous articles have addressed how to reduce diagnostic errors in medicine, from both practical and research perspectives [110–117]. Some potential components to the solution include better physician education in neurological emergencies that encourage detailed history-taking and systematic physical examination, improved access to supportive diagnostic tests (MRI), real-time neurology consultation and communicate clearly with patients and other physicians who will be seeing them in followup. Further well-designed studies are needed in the area of misdiagnosis of neurological emergencies to improve patient care and the use of healthcare resources.

References


